

## FERREED SORBENT. PREPARATION TECHNIQUE

### SPHERE OF TECHNIQUE

The inferred invention pertains to biology and medicine and might be applied for biological fluids purification and normalize condition of those to physiological standards.

### TECHNIQUE PRECEDING LEVEL

There is a known ferreed sorbent (FS), made of iron in the form of crystals with particle size of 10-15 nm (refer to e.g. certificate of authorship № 1589327, USSR, with precedence date from April 14<sup>th</sup>, 1988, IPC: C 01 G 49/08).

While exerting bactericidal effects, the known sorbent, however, is limited in applicability, as it can be used "in vitro" only.

The closest analogical prototype product is ferreed sorbent (FS), with the atomic centre (core) as grading fraction with particle size of (0,1-1000) mc, made of iron or iron oxides, or nickel, or iron-nickel alloy, and coated with one or double layer coat of carbon, or aluminum oxide, or silicium dioxide, or zirconium dioxide, or dextrane, e.g. sephadex, or gelatin, or albumin, or polysaccharide, e.g. amylum, or ion-exchange resins, e.g. cations or anions, where the coat upper layer is either conjugated with antibodies, or modified by pharmaceutical composition, e.g. antibiotics or phthalhydrazide salines, e.g. 5-amino-2,3-dihydro-1,4-dione salines, or else fermented e.g. with urease (refer to e.g. Russian Federation Patent № 2178313 with priority date from August 29<sup>th</sup>, 2000, IPC: A61M1/16).

The above sorbent appears an effective remedy used for biological fluids extracorporeal restoration to physiological standards, providing clearance of e.g. blood from low-molecular, medium-molecular and high-molecular exotoxines and endotoxines with distraction of its rheological properties; correction of biological fluids enzymatic and immune constitution, as well as antiseptis of viruses and retroviruses pathogenic microflora. However, as such sorbent turns up a very expensive product, and a great quantity of the above sorbent is needed for having an appropriate course of treatment, and consequently the treatment is related with significant financial expenses.

There is a known method of ferreed sorbent preparation technique (refer to e.g. USSR certificate of authorship № 1589327, with precedence date from April 14<sup>th</sup>, 1988, IPC: C 01 G 49/08), including iron powder volatilization procedure at low temperature ( $10^4 \times (0,5-5)^\circ \text{K}$ ) plasma in argon atmosphere, and the derived volatile product is quenched and condensed in argon gas flow, and then the precipitated product in the form of crystals is transferred to stabilizer containing dispersion medium, e.g. water at pH 7-9 or oil, and sustained there while being mixed, within (10-15) hours at the temperature (50-90) $^\circ \text{C}$  and at residual pressure of (1-3) MmHg till the end of gas liberation.

The known method ensures the possibility to derivate sorbent in the form of iron particles (crystals) with particle size of (10-15) nm, however, due to small particle size the above sorbent has got low magnetic susceptibility values, consequently in order to withdraw sorbent out of the biological medium application of magnetic fields with intensity (1-3) tesla is required, which is unacceptable by medical norms (refer to, e.g. specification to the Russian Federation Patent № 2109522 with precedence dated from August 1<sup>st</sup>, 1996, IPC: A61M1/36).

The closest analogical prototype of ferreed sorbent preparation technique (refer to e.g. Russian Federation Patent № 2109522 with precedence dated from August 1<sup>st</sup>, 1996, IPC: A61M1/36), including fractionating of high dispersed powder of Ferram reductum in inert gas flow with the velocity of (0,02-1,00) m/s under exposure of magnetic field with the intensity of  $(10-10^3) A/u$  with subsequent thermal treatment of the received iron particles at the temperature of (1000-1500)° C in inert gas flow containing coal and/or silicon oxide and/or aluminium oxide microparticles, after which treatment the ferreed sorbent particles surface are covered by biologically active compounds – food proteins or dextrane, or pharmaceutical preparations, or antibodies.

Such method ensures/provides possibility to receive ferreed sorbent of certain chemical composition, effective at recession «in vivo» and «in vitro» of low-, medum- and high molecular toxins, microflora and retroviruses. However, the above method is limited in possibility of receiving of the ferreed sorbent with volumetrical particles, having predominantly proportionate dimensions with respect to both thickness of (0,5-2,5) mcm and those particles surface dimensions corresponding to that form.

## INVENTION DISCLOSURE

The “Ferreed Sorbent” invention based on the objective to develop the sorbent similar in performance to analogous sorbent having substantially larger particles surface without any significant increase in weight of the sorbent core.

The “Ferreed Sorbent” invention based on the objective to develop the procedure providing the possibility of receiving the sorbent with the core in a form of e.g. flake.

The assigned task is achieved by enabling the fact that in the ferreed sorbent having a ferromagnetic core, with one or double layer coat or no coat, the core is made in a form of a flake, with in-plane dimensions of (500-5000) mc, and thickness of (0,1-1000) mc. Here the core is made either of iron, or nickel, or iron-nickel alloy, or iron or nickel alloy with titanium, or iron or nickel alloy with tantalum, or iron-nickel-titanium alloy, or iron-nickel-tantalum alloy.

Furthermore, the one layer coat is made either of carbon, or aluminum oxides, or silicon dioxide, or zirconium dioxide, or dextrane, e.g. from sephadex, or gelatin or albumin, or polysaccharide, e.g. amylum, or ion-exchange resins, e.g. cations or anions.

Here, in double layer coat the first closest to the core (inner) layer is made either of carbon, or aluminum oxides, or silicon dioxide, or zirconium dioxide, and the second (outer) layer of the coat is made either of dextrane, e.g. from sephadex, or

gelatin or albumin, or polysaccharide, e.g. amyllum, or ion-exchange resins, e.g. cations or anions.

Furthermore, the outer layer of the coat is either conjugated with antibodies, or modified by pharmaceutical composition, e.g. antibiotics or phthalhydrazide salines, e.g. 5-amino-2,3-dihydro-1,4-dione salines, or else fermented e.g. with urease.

The assigned task is achieved by enabling the fact that in ferreed sorbent generation method distinguishing by that iron or/and nickel or/and titanium or/and tantalum powder is volatilized or fused in a low-temperature plasma with the temperature of  $10^4 \times (0,5-5)^\circ\text{K}$ , and the received product of vaporous or fused particles of respective metals or respective metals alloys is quenched and condensed in gas flow, e.g. argon flow and then the product settled as crystals or, correspondingly, as microbars of respective metals alloys, is transferred to disperse medium containing stabilizer, e.g. water and/or oil, and while being mixed, sustained there within (5-15) hours at the temperature  $(50-90)^\circ\text{C}$  and at residual pressure of (1-5) MmHg until gas liberation ends, and after that those crystals or microbars are treated by flattening e.g. through pressing e.g. in a ball mill, until getting flakes of the specified thickness, and afterwards are repeatedly (up to 10 times) washed in distilled water, and then separated from weak parts of flakes, treating with e.g. ultrasound of e.g. (200-300)  $\text{Vt/cm}^2$  capacity, and then the received flakes dried out e.g. in a hot air sterilizer at the temperature of  $(80-110)^\circ\text{C}$ , and after that the dried flakes are fractionated in either inert gas flow with the velocity of (0,02-1,00) m/s under exposure of magnetic field of  $5 \times (10-10^3)$  A/m intensity, or by using e.g. centrifugation. Then the specified size sorbent cores with layer-by-layer formed coat are extracted, and the received end product is packed in light-protected and hermetically sealed containers and sterilized, by e.g. U-rays, where sorbent received right after fractionating can be used as the end product as well.

Here the first (inner) layer of the coat is formed by thermal treatment of fractionated flakes at the temperature of  $(1000-1500)^\circ\text{C}$  in inert gas flow, e.g. flow of argon, containing microparticles of either carbon, or silicon oxide, or aluminum oxide, or zirconium oxide.

Furthermore, the first layer of the coat is formed by blending with using ultrasound exposure to fractionated flakes suspension within (1-10) minutes in heated to the temperature of  $(30-80)^\circ\text{C}$  aqueous solution of dextrane, or gelatin or albumin, or amyllum, with subsequent cooling of the above suspension down to the temperature of  $(4-10)^\circ\text{C}$ , and the received precipitate is filled up with formalin, sustained there within (10-40) minutes, simultaneously being mixed, and after that dried out thoroughly at the temperature of  $(25-50)^\circ\text{C}$  and grinded, then the received sorbent capsules (the end product) are filtered in magnetic field.

Furthermore, the first layer of the coat is formed through adding an ion-exchange resin, e.g. amberlite into fractionated flakes suspension in distilled water, heated up to the temperature of  $(40-60)^\circ\text{C}$ , with subsequent cooling of the above suspension down to the temperature of  $(15-30)^\circ\text{C}$ , with adding nitrous acid ( $\text{HNO}_2$ ) diluted in water, sustaining within (10-15) minutes, cooling down to the temperature of  $(4-10)^\circ\text{C}$  and elution of precipitate which is washed in physiological solution and buffered in aqueous solution of  $\text{NH}_4\text{OH}$  foundation blend and  $\text{NH}_4\text{Cl}$  salt.

Here the second layer of the coat is formed by blending with using ultrasound

exposure within (1-10) minutes to suspension of ferromagnetics covered with carbon or silicon oxide, or aluminium oxide, or zirconium oxide coat in aqueous solution of dextrane, or gelatin, or albumin, or amylum heated up to the temperature (30-80)° C with subsequent cooling of the above suspension down to the temperature (4-10)° C, and the received precipitate is filled up with formalin, sustained in there within (10-40) minutes simultaneously being mixed, then dried out thoroughly at the temperature of (25-50)° C, grinded and the received sorbent capsules (of the end product) are filtered in magnetic field.

Furthermore, the second layer of the coat is formed through adding an ion-exchange resin, e.g. amberlite into heated up to the temperature of (40-60)° C suspension of ferromagnetics, covered with carbon or silicon oxide, or aluminium oxide, or zirconium oxide coat, in distilled water, with subsequent cooling of the above suspension down to the temperature of (15-30)° C, and adding and immixturing albumin, e.g. in the form of serum, with subsequent adding of nitrous acid (HNO<sub>2</sub>) diluted in water, sustaining within (10-15) minutes, cooling down to the temperature of (4-10)° C and elution of precipitate which is activated by sustaining within (1,5-2) hours in a modifier solution, then washed in physiological solution and buffered in aqueous solution of NH<sub>4</sub> OH foundation blend and NH<sub>4</sub> Cl salt.

Sodium periodate (NaIO<sub>4</sub>) or glutaric dialdehyde in (3-10)% solution of Na<sub>2</sub>SO<sub>4</sub> in water can be used here as a modifier.

Furthermore, while forming the outer layer of the coat, it is conjugated with antibodies by adding the ferreed sorbent with one or double layer coat into aqueous suspension; but it should be with the outer coat layer made of sephadex or albumin, modified e.g. with glutaric dialdehyde or sodium periodate, with serum, e.g. of blood, containing antibodies, specified to sorbed antigen, e.g. to systemic lupus erythematosus antigen, in buffered liquid with pH of 6,5-10, sustaining while being mixed of the above compound within (1-3) hours at the temperature of (15-25)° C, subsequent adding to the compound of sodium borhydrate, cooling down to the temperature of (4-10)° C, repeated sustaining while being mixed within (1-3) hours, precipitate extraction and its buffering and drying out.

Furthermore, while forming the outer layer of the coat, it is modified with pharmaceutical composition through heating the ferreed sorbent suspension with one or double layer coat (but with the outer coat made of e.g. dextrane or gelatin) up to the temperature of (35-70)° C in physiological solution, adding into there a pharmaceutical composition in powder, e.g. antibiotic, e.g. oxaccillin, sustaining at thorough mixing at the above mentioned temperature within (0,5-2,5) hours, subsequent cooling of the above compound down to the temperature of (4-10)° C, decanting of supernatant fluid in magnetic field, washing the precipitate in running distilled water and its subsequent drying out.

Furthermore, while forming the outer layer of the coat, it is modified through preliminary dissolution of urease crystals in polyether, e.g. dibenzo-18 crown 6, immixture of the above solution with the suspension in distilled water of ferreed sorbent with the coat made of e.g. sephadex, sustaining while being mixed at the temperature of (25-40)° C within (2-5) hours and cooling down to the temperature of (4-10)° C, subsequent adding of formaldehyde and repeated sustaining within (1-3) hours, draining out the supernatant fluid in the presence of magnetic field and drying out the precipitate.

Furthermore, while forming the outer layer of the coat, it is modified through heating up aqueous suspension of the ferreed sorbent with the coat made of e.g. dextrane, to the temperature of  $(40-70)^{\circ}\text{C}$ , subsequent immixture with zirconium saline powder, e.g. of respective phthalhydrazide saline, and  $(50-120)\text{ Vt/cm}$  intensity ultrasound exposure to the above mixture within  $(1-10)$  minutes, cooling the received compound down to the temperature of  $(4-10)^{\circ}\text{C}$ , adding formaldehyde, sustaining while being mixed within  $(1-3)$  hours, draining out the supernatant fluid in the presence of magnetic field and drying out the precipitate.

### THE BEST VARIATION OF THE INVENTION EMBODIMENT

Ferreed sorbent is made in the form of cores with one or double layer coat surrounding the core; and with no coating.

To be used as cores for the ferreed sorbent they take powder from ferromagnets, e.g. from iron (Fe), its oxides ( $\text{Fe}_2\text{O}_3$  or  $\text{Fe}_3\text{O}_4$ ), nickel (Ni), iron-nickel alloys, as well as from iron or nickel alloy with titanium (Ti), or from iron or nickel alloy with tantalum (Ta), from iron-nickel-titanium alloy, or from iron-nickel-tantalum-titanium alloy and the like magnetic sensible materials.

For the subsequent use fractions in the form of flakes with the dimensions in plane of  $(500-5000)\text{ mc}$  and with the thickness of  $(0,1-1000)\text{ mc}$  are taken.

For getting cores for the ferreed sorbent, iron, or/and nickel, or/and titanium, or/and tantalum powder with particle size of  $(10^2-10^5)\text{ nm}$  is volatilized or/and fused in low-temperature plasma with the temperature of  $10^4 \times (0,5-5)^{\circ}\text{K}$ , and the received product volatilized or/and fused in the form of respective metals or respective metals alloys with concentration of  $(0,1-0,5)$  volume % quenched down to the temperature of  $(50-80)^{\circ}\text{C}$  and condensed in reactor (refer to e.g. description to USSR certificate of authorship № 1589327 prioritized date from April 14<sup>th</sup>, 1988, IPC: C 01 G 49/08) in gas flow, e.g. in argon flow, and then the product settled in the form of crystals or, respectively, microbars of respective metals alloys, e.g. in the amount of  $(0,05-10)\text{ mg}$ , is transferred to the disperse medium containing stabilizer, e.g. distilled water of  $(50-500)\text{ ml}$  with pH of 7-9 and/or mineral, e.g. paraffin or vegetable oil e.g. olive or sea-buckthorn oil, with preliminarily added e.g. oleic acid in the amount of  $(2-20)$  volume %, and, while being mixed, sustained in there within  $(5-15)$  hours at the temperature of  $(50-90)^{\circ}\text{C}$  and at the residual pressure of  $(1-5)\text{ Mmmhg}$  till the end of gas liberation.

After that those crystals or microbars are treated by flatter, e.g. through pressing e.g. in a ball mill, till having flakes of the specified thickness, which then repeatedly (up to 10 times) are washed in distilled water, and then weak flake parts are removed by exposing to ultrasound of e.g.  $(200-300)\text{ Vt/sm}^2$  intensity in e.g. water.

The received material (different size flakes and chip bits) is dried in whole e.g. in a hot air sterilizer at the temperature of  $(80-110)^{\circ}\text{C}$ , and then the dried product (flakes) fractionated either in inert gas flow with velocity of  $(0,02-1,00)\text{ m/c}$  under the exposure of magnetic field with intensity of  $5 \times (10-10^3)\text{ A/m}$  or by using centrifugation, and the sorbent (flakes) of the specified size is excreted in the form of cores, on which coats are formed layer by layer, and the acquired end product is packed up in lightproof hermetically closed containers and sterilized through e.g. U-

rays; here the sorbent received right after fractionating can be chosen as the end product as well. The output of conditioned sorbent cores after fractionating makes (60-75) %.

For getting (forming) of the first (closest to the core) layer of the coat, the fractionated flakes are treated (at the temperature of (1000-1500)° C) in thermo oven in inert gas flow, e.g. in argon flow, containing microparticles of carbon (C), or silicon oxide (SiO<sub>2</sub>), or aluminum oxide (Al<sub>2</sub>O<sub>3</sub> or Al<sub>3</sub>O<sub>4</sub>), or zirconium oxide (ZrO<sub>2</sub>). Flow velocity makes (0,02-1,2) m/s. Coating quality of cores depends on inert gas flow throughput rate, as well as on saturation of the gas with microparticles of coating material and the size of those particles. In the given examples the thickness of the coat layer made with the above method makes (0,2-50) mc.

Efficient output of sorbent - (70-85) %.

While forming of the first layer of the coat through covering sorbent cores with such substances like either dextrane, or gelatin, or albumin, or amylum, fractionated flakes suspension in the amount of (2-20) g in (10-50) ml of distilled water is mixed with (50-100) ml of heated to the temperature of (30-80)°C aqueous solution of either dextrane, or gelatin, or albumin, or amylum, with the blend ratio of (volume %): (50-95) % -respective product, the rest is water; then mixed within (1-10) minutes till it gets homogeneous structure under the exposure of e.g. ultrasound dispergator "УЗДН-2Т" (refer to e.g. the description to the certificate of authorship № 1684616, USSR) ultrasound with oscillation frequency (10-15) kHz and intensity rate of (50-120) Wt/cm; then the suspension is cooled e.g. in a refrigerator down to the temperature of (4-10)°C, then the precipitate received is filled up with formalin (aqueous solution HCHO), sustained in there within (10-40) minutes while simultaneously being mixed, after that it should be thoroughly dried up at the temperature of (25-50)° C, grinded and the received sorbent capsules (the end product) are filtered in magnetic field with the intensity of  $5 \times (10^{-10})^3$  A/m, of e.g. constant magnet made of samarium (8t)-cobalt (Co) alloy.

Thickness of the coat layer made using the method above makes (0,5- 3) mm.

Quantitative output of sorbent makes (85-95) % out of initial.

While forming of the first layer of the coat by using ion-exchange resin, e.g. (10-25) g of amberlite is added into heated up to the temperature of (40-60)° C fractionated flakes suspension of (2-5) g per (10-100) ml of distilled water, then the received compound is cooled down to the temperature of (15-30) ° C, add nitrous acid (HNO<sub>3</sub>) diluted in water (in the amount of (1-10) vol. %), sustained within (10-15) minutes, then cool again down to the temperature of (4-10)°C and then precipitate is excreted, which is washed in physiological solution, buffered till it gets pH 4,0±0,5 in aqueous solution of foundation of NH<sub>4</sub>OH or NH<sub>4</sub>Cl saline.

Thickness of the coat layer made by the above method makes (0,2- 1) mm.

Quantitative output of sorbent makes (90-92) % out of initial.

While forming of the second layer of the coat through covering the ferreed sorbent coated with either carbon or silicon oxide or aluminum oxide, or zirconium oxide with such substances like either dextrane, or gelatin, or albumin, or amylum, a suspension of ferromagnetics (in the amount of (2-20) g per (10-50) ml of distilled water), covered with carbon, or silicon oxide, or aluminum oxide, or zirconium oxide coat, being mixed within (1-10) minutes under the exposure of ultrasound with intensity of (50-120) Vt/cm in (50-100) ml of heated to the temperature of (30-80)°

C (50-95) % solution of dextrane, or gelatin, or albumin, or amylum in distilled water with subsequent cooling to the above suspension down to the temperature (4-10)° C. The precipitate is filled up with formalin, sustained in there within (10-40) minutes while simultaneously being mixed and after that it is thoroughly dried out at the temperature of (25-50)° C, grinded, and the acquired sorbent capsules (end product) are filtered in magnetic field with the intensity of  $5 \times (10-10^3)$  A/m.

Thickness of the coat layer made by the above method makes (0,5- 3) mm.

Quantitative output of sorbent makes (85-95) % out of initial.

While forming of the second layer of the coat by using ion-exchange resin, a suspension of ferromagnetics (in the amount of (0,2-0,5) g per (10-100) ml of distilled water), covered with carbon, or silicon oxide, or aluminum oxide, or zirconium oxide coat, heated up to the temperature of (40-60)° C, then e.g (1-2) g of amberlite is added into there, and then the received compound is cooled down to the temperature of (15-30) ° C, then nitrous acid (HNO<sub>3</sub>) diluted in water (in the amount of (1-10) vol. %) is added, sustained within (10-15) minutes, then cooled again down to the temperature (4-10)° C and the precipitate is excreted, which is activated by sustaining within (1,5-2) hours in a modifier solution, then washed in physiological solution and buffered till it gets to pH 4,0±0,5 in aqueous solution of NH<sub>4</sub>OH foundation and NH<sub>4</sub>Cl salt. Here sodium periodate (NaIO<sub>4</sub>) or glutaric dialdehyde in (3-10)% solution of Na<sub>2</sub>S04 in water can be used as modifier.

Thickness of the coat layer made by the above method makes (0,2- 1) mm.

Quantitative output of sorbent makes (90-95) % out of initial.

Moreover, while forming the outer layer of the coat, it is conjugated with antibodies through adding serum e.g. of blood, into aqueous suspension of ferreed sorbent with one or double coated, but with the outer coat made from sephadex or albumin, modified with e.g. glutaric dialdehyde or sodium periodate, (in the amount of (1-50) ml of serum per (100-150) ml of suspension), containing antibodies specific to the antigen sorbed, e.g. to systemic lupus erythematosus antigen, in buffering liquid with pH of 6,5-10, sustaining while the compound being mixed within (1-3) hours at the temperature of (15-25)° C, subsequent adding of sodium borhydrate into the compound, cooling down to the temperature of (4-10)° C, repeated sustaining with simultaneous mixing within (1-3) hours, precipitate extraction and its buffering and drying out.

Here the respective coat layer thickness is increased for (0,2- 0,5) mm.

Quantitative output of sorbent makes (92-95) % out of initial.

Furthermore, while forming of the outer layer of the coat, it is modified with pharmaceutical composition by heating up to the temperature of (35-70)° C of aqueous suspension of ferreed sorbent ((10-20) g of sorbent per (50) ml of distilled water) with one or double layer coat, but the outer coat made of e.g. dextrane, or gelatin, in physiological solution (0,9 % solution of NaCl in distilled water), adding a pharmaceutical preparation powder (in the amount of (1-5) r per (10-50) ml of suspension), e.g. antibiotic, e.g. oxaccillin, sustaining while simultaneous thorough mixing at the above mentioned temperature within (0,5-2,5) hours, subsequent cooling of the above compound down to the temperature of (4-10)° C, decanting of the supernatant fluid in magnetic field with the intensity of  $5 \times (10-10^3)$  A/m, washing the precipitate in running distilled water and its subsequent drying out at the temperature of (25-40)° C.

Here the respective coat layer thickness is increased for (0,01- 0,1) mm.

Quantitative output of sorbent makes (90-95) % out of initial.

Furthermore, while forming of the outer layer of the coat, it is modified by preliminary dilution of e.g. (1-5) g of urease crystals in (10-15) ml of polyether, e.g. of dibenzo-18 crown 6, blending the above solution with ferreed sorbent suspension in distilled water ((10-15) hg of sorbent per (50-100) ml of water) with the coat made e.g. from sephadex-10, sustaining while mixed at the temperature of (25-40)° C within (2-5) hours and cooling down to the temperature of (4-10)° C, subsequent adding of formaldehyde ((25-30) ml per 100 ml of compound) and repeated sustaining while mixed within (1-3) hours, pouring out the supernatant fluid under the influence of magnetic field with the intensity of  $5 \times (10-10^3)$  A/m and precipitate drying out e.g. in a hot air sterilizer at the temperature of (50-85)° C.

Here the respective coat layer thickness is increased for (0,5- 1) mm.

Quantitative output of sorbent makes (90-95) % out of initial.

Furthermore, while forming of the outer layer of the coat, it is modified through heating of aqueous suspension of ferreed sorbent with e.g. dextrane coat up ((15-20) g of sorbent per 75-100 ml of distilled water) to the temperature of (40-70)° C, subsequent blending with zirconium saline powder of e.g. respective phthalhydrazide saline, e.g. 5-amino-2,3-dihydro-1,4-dion, and treating the above compound within (1-10) minutes with ultrasound of (15-25) kHz oscillation frequency and (50-120) Vt/cm<sup>2</sup> intensity, cooling of the received compound down to the temperature (4-10)° C, adding formaldehyde ((25-30) ml per 100 ml of compound), sustaining in there while mixing within (1-3) hours, pouring out of supernatant fluid in the presence of magnetic field with the intensity of  $5 \times (10-10^3)$  A/m and precipitate drying out at the temperature of (25-45) C.

Here the respective coat layer thickness is increased for (0,01- 0,1) mm.

Quantitative output of sorbent makes (90-95) % out of initial.

### INDUSTRIAL APPLICABILITY

Use of ferreed sorbent having substantially larger surface of the particles with no significant weight increase of its core, and the method of receiving such sorbent allows to provide effective cleaning of biological fluids, e.g. blood, out of low-, medium- and high-molecular exotoxines and endotoxines without disorder of its rheological properties, provide possibility to correct ferment and immune structure of the biological fluids, as well as destruction of viruses and retroviruses pathogenic microflora while using appreciably low amount of the proposed ferreed sorbent (with respect to weight) relatively to the amount of the analogous sorbent known earlier and specified for the same purposes.

Thus, in the view of the fact that biological fluid cleaning by using ferreed sorbent is taken place by interaction of its surface with the fluid being corrected, one can show that the effective particle surface of the known sorbent, size of which in terms of length, width and thickness are on average commensurable at mass conservation, is significantly smaller than the surface of the proposed sorbent.

For example let's take a spherical particle.

Using known mathematical formulas, we will get the following (as sphere volume value ( $V_{\text{sphere}}$ ) is equal to:



$V_{\text{sphere}} = 4\pi r^3/3$ , where  $r$  – sphere radius, and accordingly, the sphere surface area ( $S_{\text{sphere}}$ ) is equal to  
 $S_{\text{sphere}} = 4\pi r^2$ , then  
 $S_{\text{sphere}} = 3 V_{\text{sphere}} / r$

Considering that particle mass is proportional to its volume, and assuming that after above described procedure of acquiring sorbent particles in the form of flakes, a spherical sorbent particle will be reformed into round flake/disk, then as the flake volume is  $V_{\text{flake}} = \pi R^2 \delta$ , and the surface area  $S_{\text{flake}} = \pi R^2$ , where  $R$  – flake radius, and  $\delta$  – its thickness, while  $\delta = 0,1$  g (in accordance with the abovesaid statement about some decrease of particle thickness), then we have  $S_{\text{flake}} = V_{\text{flake}} / 0,1$  g.

Considering that  $V_{\text{flake}} = V_{\text{sphere}}$ , then, as their masses are equal, we will get the following:

$$S_{\text{flake}} = 10 V_{\text{sphere}} / r$$

Taking into consideration that, there are two such surfaces on the flake, and putting the term (1) into the formula (2) we will get the following:

$$S_{\text{flake's full surface}} = 20 S_{\text{sphere}} / 3$$

The results received justify the above hypothesis that in case of using of the sorbent being proposed the each particle surface interacting with biological fluid is significantly enlarged, and, consequently, consumption of sorbent and respective treatment costs are decreased.

Feasibility of effective application of the proposed ferreed sorbent extracted using the above-described methods is testified by the following examples:

Example 1. A non-pedigree dog weighing 12 kilos was injected (per os) 4,3 g of veronal. After 45 minutes amount of barbiturate in blood gets 118 mkg/ml.

Blood extracorporeal regeneration (correction) procedure was conducted using the expedient equipment (УКБЖ-1). The animal's blood was retrieved in portions of 10 ml, being then blended in equal volume proportions with ferreed sorbent suspension in physiological solution, which contained (mass. %): ferreed sorbent (core – nickel flake, coat inner layer - carbon, coat outer layer - dextrane) - 1,5; anticoagulant (heparin) -0,015; physiological solution as the balance; then the blood was sustained within 2-3 seconds and administered back into animal's organism.

About one liter of blood had been treated/processed during one session.

Indications before and after the correction session:

Creatinine (m mole/l)	1,45	1,10.
Urea (m mole /l)	11,9	6,2.
Bilirubin (total) (m mole /l)	25,0	14, 4.
Barbiturates (mkg/ml)	141,5	14,2.

Furthermore, gastric lavage was made during the session, the animal was injected intravenously 500 ml of solution of electrolytes and 2 % glucose.

After the session, the animal was in the state of moderate severity, brisk reflexes.

The indications of sorbate effectiveness are shown in the following examples

below, as well as effectiveness of selective and functional properties of known ferreed sorbents, described, e.g. in the specifications to the patent of Russian Federation № 2178313 (refer to e.g. Russian Federation patent 2178313 with the priority dated from 29.08.2000, IASC: A 61 M 1/16) and the results received during the researches with ferreed sorbent being proposed in the present invention hereby.

Example 2. 5 ml of carbofos solution was injected into the test-tube with 100 ml of a non-pedigree dog blood. Carbofos concentration in the blood was 0,015 mkg/ml.

The received blend was divided in two parts and each part was added 20 ml of ferreed sorbent suspension, where one part was added the known ferreed sorbent suspension in physiological solution (cores as iron particles, coat layers as silicon oxide) in the amount of 1,0 g, while the second part was added the proposed ferreed sorbent with the same material composition but with flake cores, in the amount of 0,01 g.

After mixing of the received compositions within 1,5 minutes the supernatant fluid was decanted, and the precipitate was withheld using a magnet.

Carbofos concentration in the supernatant fluid received from the first blend made 0,002 mkg/ml, and the supernatant fluid received from the second blend made 0,012 mkg/ml.

Example 3. Into two different test-tubes each containing 20 ml of blood serum of a dog with simulated nephratonia (urea concentration in the first test-tube was 26,4 m mole/l, and 30,2 m mole/l in the second), the following had been added: 200 mg of the known ferreed sorbent with the coating of sephadex-10 fermented with urease into the first test-tube; 30 mg of the ferreed sorbent being proposed with the cores in the form of titanium flakes with the coating analogous to the above specified – into the second test-tube.

After sustaining (while shake) of the received compositions within 5 seconds and removal of the supernatant fluid in magnetic field, the urea content concentration in supernatant fluid in the first test-tube got - 10,7 m mole/l, and it got 12,1 m mole/l in the second one.

Example 4. In two different test-tubes each containing 20 ml of phosphoric acid sodium saline solution ( $\text{NaH}_2\text{PO}_4$ ) in water the following had been added: 100 mg of the known ferreed sorbent with cation-modified (COON group polysaccharides) ion-exchange resin coating into the first test-tube; 10 mg of the ferreed sorbent being proposed in the form of tantalum flakes with the coating analogous to the above specified - into the second test-tube.

After mixing (while shake) of the received compositions and removal of the supernatant fluid in magnetic field, the concentration of phosphates in the supernatant fluid received from the first test-tube had reduced for 57 % comparatively to the initial, and the concentration of phosphates in the supernatant fluid from the second test-tube, correspondingly, had reduced for almost half (for 44,8 %) from the initial point of phosphates concentration.

Example 5. In two different test-tubes each containing 20 ml of sulphuric acid salines solution in water the following had been added: 100 mg of the known ferreed sorbent with anionite-modified ( $\text{NH}_3^+$  group) ion-exchange resin coating into the first test-tube; 20 mg of the ferreed sorbent being proposed in the form of iron-nickel flakes with the coating analogous to the above specified – into the second test-tube.

After mixing (while shake) of the received compositions and removal of the supernatant fluid in magnetic field, the concentration of sulphuric acid salines in the supernatant fluids received from both of the test-tubes had reduced virtually for the same i.e. for 72 % comparatively to the initial concentration in the first test-tube, and for 73, 4 % comparatively to the initial concentration – in the second test-tube.

Example 6. In two different test-tubes each containing 20 ml of blood of a patient with chronic renal-hepatic insufficiency disease the following had been added: 100 mg of the known ferreed sorbent with zirconium luminole saline-modified dextrane coating into the first test-tube; 30 mg of the ferreed sorbent being proposed in the form of iron-titanium flakes with the coating analogous to the above-specified – into the second test-tube.

After mixing (while shake) of the received compositions and removal of the supernatant fluid in magnetic field, the concentration of phosphoric acid salines ( $\text{NaH}_2\text{PO}_4$ ) in the supernatant fluid received from the first test-tube had got 0,07 mg/ml; and the concentration of phosphoric acid salines ( $\text{NaH}_2\text{PC}^{\wedge}$ ) in the supernatant fluid received from the second test-tube had got 0,021 mg/ml. (The initial concentration of the saline was 0,61 mg/ml).

Example 7. In two different test-tubes each containing 10 ml of blood serum of a patient with chronic renal-hepatic insufficiency disease the following had been added: 50 mg of the known ferreed sorbent with iron-nickel cores and urease-modified sephadex coating into the first test-tube; 10 mg of the ferreed sorbent being proposed with iron-nickel cores with coating analogous to the above-specified – into the second test-tube.

After sustaining within 10 seconds and the supernatant fluid decanting (sorption) the urea concentration in the supernatant fluid received from the first test-tube had reduced for 23 % comparatively to the initial urea concentration in blood serum, and the urea concentration in the supernatant fluid received from the second test-tube had reduced for 35 % comparatively to the initial urea concentration in blood serum.

Example 8. In two different test-tubes each containing 20 ml of blood serum of a patient with sepsis the following had been added: 150 mg of the known ferreed sorbent with iron-nickel cores and oxaccillin-modified gelatin coating into the first test-tube; 15 mg of the ferreed sorbent being proposed with iron-nickel-titanium-tantalum alloy flake cores with coating analogous to the above-specified – into the second test-tube.

After mixing while shaking of the test-tubes contents within 2 minutes, the supernatant fluid was decanted and the hard constituent was retained using a magnet field.

Inoculation was made both on the patient's blood agar-agar and the blood having been exposed to ferreed sorbent (the supernatant fluids) from the both test-tubes.

Growth of streptococcus and staphylococcus colonies was observed in the inoculation of the patients' blood; no such growth was observed in the inoculation of the blood taken from the test-tubes.

Example 9. In two different test-tubes each containing 10 ml of lymph plasma of a patient with sepsis the following had been added: 100 mg of the known ferreed sorbent with iron-nickel cores and dextrane coating into the first test-tube; 15 mg of

the ferreed sorbent being proposed with iron-nickel-titanium-tantalum alloy flake cores with coating analogous to the above-specified – into the second test-tube.

After mixing (while shaking) of the compositions received and removal of the supernatant fluid in magnetic field, inoculation was made both on the patient's lymph agar-agar and the lymph having been exposed to ferreed sorbent (the supernatant fluids) from the both test-tubes.

Growth of multiple staphylococcus colonies was observed in the inoculation of the lymph with no lymph-separation; virtually no such growth was observed in the inoculation of the supernatant fluids taken from the test-tubes.

Example 10. In two different test-tubes each containing 5 ml of blood-tinted cerebrospinal fluid (a patient with craniocerebral injury) the following had been added: 50 mg of the known ferreed sorbent with iron cores and silicon oxide coating into the first test-tube; 15 mg of the ferreed sorbent being proposed with iron-tantalum alloy flake cores with coating analogous to the above-specified – into the second test-tube.

After sedimentation the cerebrospinal fluid in the test-tubes had got light yellow colour.

Effectiveness of the developed preparation application is testified in the experiments when doing the research on sorption capacity of the ferreed sorbent for each above-described variation for its performance, and at the same time the results commensurable to the results of using analogous variations of the known ferreed sorbent were achieved at using significantly lower amounts of the ferreed sorbent being proposed.

## INVENTION FORMULA

1. Ferreed sorbent involving a ferromagnetic core, with one or double layer coat or with no coat, distinguished by that the core is made in the form of a flake, with the dimensions in plane of (500-5000) mc and with the thickness of (0,1-1000) mc.

2. Ferreed sorbent as per paragraph 1 distinguished by that the core is made either of iron, or iron oxides, or nickel, or iron-nickel alloy, or iron- or nickel – titanium alloy, or iron- or nickel-tantalum alloy, or iron-nickel-titanium alloy, or iron-nickel-tantalum alloy.

3. Ferreed sorbent as per any of paragraphs 1-2 distinguished by that the one-layer coat is made of either carbon, or aluminium oxides, or disilcon oxide, or zirconium dioxide, or dextrane, e.g. of sephadex, or gelatin, or albumin, or polysaccharide, e.g. of amylum, or ion-exchange resins, e.g. cations or anions.

4. Ferreed sorbent as per any of paragraphs 1-2 distinguished by that in the double-layer coat the first closest to the core (inner) layer is made of either carbon, or aluminium oxides, or disilcon oxide, or zirconium dioxide, and the second (outer) layer of the coat is made either of or dextrane, e.g. of sephadex, or gelatin, or albumin, or polysaccharide, e.g. of amylum, or ion-exchange resins, e.g. cations or anions.

5. Ferreed sorbent as per any of paragraphs 1, 3-4 distinguished by that the outer layer of the coat is made by either conjugation with antibodies or modification with a pharmaceutical preparation, e.g. antibiotics or phthalhydrazide salines, e.g 5-

amino-2,3-dihydro-1,4-dione salines, or made fermented e.g. with urease.

6. Method of extraction of ferreed sorbent as per any of paragraphs 1-2, distinguished by that the powder of either iron or/and nickel, or/and titanium, or/and tantalum is volatilized or/and fused in low-temperature plasma with the temperature  $10^4 \times (0,5-5)^\circ\text{K}$ , and the received vaporous or/and being in the form of fused particles of the respected metals or respected metals alloys product is quenched and condensed in a gas flow e.g. argon flow, and then the product precipitated in the form of crystals or, correspondingly, as respected metals alloys microbars, is transferred into the dispersion medium containing stabilizer. E.g. water and/or oil, and while being mixed, it is sustained there within (5-15) hours at the temperature of  $(50-90)^\circ\text{C}$  and at the residual pressure of (1-5) MmHg till the end of gas liberation, after that the crystals or microbars are treated by flattening e.g. by pressing, till they get to flakes of the specified thickness, which then repeatedly (up to 10 times) are washed in distilled water and then flakes weak parts are removed by exposing to ultrasound (e.g. in water) of e.g.  $(200-300) \text{ Vt/cm}^2$  intensity, then the received flakes are dried out e.g. in a hot air sterilizer at the temperature of  $(80-110)^\circ\text{C}$ , after that the dried flakes are fractionated in either inert gas flow with the velocity of  $(0,02-1,00) \text{ m/s}$  at the exposure of magnetic field with the intensity of  $(10-10^3) \text{ A/m}$  or using e.g. centrifugation, and then the sorbent cores of a specified dimensions are educed, on which coats are formed layer-by-layer, and the received end product is packed in light-proof hermetically sealed containers, sterilized by e.g. U-rays, at that the sorbent received right after fractionation can be chosen as the end product as well.

7. Method of extraction of ferreed sorbent as per paragraph 6, distinguished by that the first closest to the core layer of the coat is formed by thermal treatment of the fractionated flakes at the temperature of  $(1000-1500)^\circ\text{C}$  in inert gas flow of e.g. argon containing microparticles of carbon, or silicon oxide, or aluminium oxide, or zirconium oxide.

8. Method of extraction of ferreed sorbent as per paragraph 6, distinguished by that the first layer of the coat is formed though mixing with using ultrasound exposure to fractionated flakes suspension within (1-10) minutes in heated aqueous solution of dextrane or gelatin, or albumin, or amylum up to the temperature  $(30-80)^\circ\text{C}$  with subsequent quenching of the above suspension down to the temperature of  $(4-10)^\circ\text{C}$ , and the received precipitate is filled up with formalin, sustained in there within (10-40) minutes while simultaneously mixing, after that it is thoroughly dried out at the temperature of  $(25-50)^\circ\text{C}$ , grinded, and the received sorbent capsules (end product) are filtered in magnetic field.

9. Method as per paragraph 6, distinguished by that the first layer of the coat is formed though adding ion-exchange resin e.g. of amberlite into suspension of fractionated flakes in distilled water heated up to the temperature of  $(40-60)^\circ\text{C}$ , with subsequent cooling of the above suspension down to the temperature of  $(15-30)^\circ\text{C}$ , and adding in there nitrous acid ( $\text{HNO}_2$ ) diluted in water, sustaining within (10-15) minutes, cooling down to the temperature of  $(4-10)^\circ\text{C}$  and extracting of precipitate which then is washed in physiological solution, buffered in aqueous solution of a blend of  $\text{NH}_4\text{OH}$  foundation and  $\text{NH}_4\text{Cl}$  saline.

10. Method per paragraph 6, distinguished by that the second layer of the coat is formed though mixing with using ultrasound exposure within (1-10) minutes to ferromagnetics suspension covered with carbon or silicon oxide, or aluminium oxide,

or zirconium oxide coating in heated up to the temperature of (30-80)° C aqueous solution of dextrane, or gelatin, or albumin, or amylum with subsequent cooling of the above suspension down to the temperature of (4-10)° C, and the received precipitate is filled up with formalin, sustained in there within (10-40) minutes while simultaneously being mixed, and after that it is thoroughly dried out at the temperature of (25-50)° C, grinded and the received sorbent capsules (the end product) are filtered in magnetic field.

11. Method per paragraph 6, distinguished by that the second layer of the coat is formed though adding ion-exchange resin e.g. of amberlite into suspension in distilled water of ferromagnetics covered with carbon or silicon oxide, or aluminium oxide, or zirconium oxide coating, heated up to the temperature of (40-60) ° C, with subsequent cooling of the above suspension down to the temperature of (15-30)° C, and adding, while it is being mixed, albumin e.g. in the form of serum with subsequent adding in there of nitrous acid (HN0<sub>2</sub>) diluted in water, sustaining within (10-15) minutes, cooling down to the temperature of (4-10)° C and extracting of precipitate which then is activated by sustaining within (1,5-2) hours in a modifier solution, washed in a physiological solution, buffered till it gets to pH 4,0 0,5 in aqueous solution of a blend of NH<sub>4</sub> OH foundation and NH<sub>4</sub>Cl saline.

12. Method per paragraph 11, distinguished by that sodium periodate (NaIO<sub>4</sub>) or glutaric dialdehyde in (3-10)% aqueous solution of Na<sub>2</sub>SO<sub>4</sub> can be used as a modifier.

13. Method as per any of paragraphs 6-11, distinguished by that while forming of the outer layer of the coat it is conjugated with antibodies through adding into the aqueous suspension of ferreed sorbent with one or double layer coating (but with the outer layer made of sephadex or albumin, and modified with e.g. glutaric dialdehyde or sodium periodate) , of serum, e.g. of blood containing antibodies specific to the antigen being sorbed e.g. to systemic lupus erythematosus antigen, in a buffered fluid with pH of 6,5-10, further sustaining of the above composition while it is being mixed within (1-3) hours at the temperature of (15-25)° C, subsequent adding of sodium borhydrate into the composition, cooling down to the temperature of (4-10)° C, repeated sustaining while mixing within (1-3) hours, extraction of the precipitate and its buffering and drying out.

14. Method as per any of paragraphs 6-11, distinguished by that while forming of the outer layer of the coat it is modified by pharmaceutical preparation through heating the suspension of ferreed sorbent with one or double layer coating (but with the outer layer made of e.g. dextrane or gelatin) heated up to the temperature of (35-70)° C in physiological solution, adding in there of a pharmaceutical preparation powder, e.g. of antibiotics, e.g. oxaccillin, sustaining while thoroughly being mixed at the above temperature within (0,5-2,5) hours, consequent cooling of the compound down to the temperature (4-10)° C, decanting of the supernatant fluid in magnetic field, washing of the precipitate in running distilled water and its consequent drying out.

15. Method as per any of paragraphs 6-11, distinguished by that while forming of the outer layer of the coat it is modified through preliminary dilution of urease crystals in polyether, e.g. in dibenzo-18 crown 6, immixture of the above composition with suspension in distilled water of the ferreed sorbent with coating made e.g. of sephadex, consequent sustaining while being mixed at the temperature

of (25-40)° C within (2-5) hours and cooling down to the temperature of (4-10)° C, consequent adding of formaldehyde and repeated sustaining within (1-3) hours, removal of supernatant fluid under exposure of magnetic field and precipitate drying out.

16. Method as per any of paragraphs 6-11, distinguished by that while forming of the outer layer of the coat it is modified through heating of aqueous suspension of ferreed sorbent with coating made of e.g. dextrane, up to the temperature of (40-70)° C, with subsequent immixturing with zirconium saline powder, e.g. of respective phthalhydrazide saline, and (50-120) Vt/cm intensity ultrasound exposure within (1-10) minutes, cooling of the received blend down to the temperature of (4-10)° C, adding formaldehyde, sustaining while it is being mixed within (1-3) hours, removal of supernatant fluid under the exposure of magnetic field and precipitate drying out.